SEQUENCE LISTING

<110> CANFIELD, William

<120> METHODS OF PRODUCING HIGH MANNOSE GLYCOPROTEINS IN COMPLEX CARBOHYDRATE DEFICIENT CELLS

<130> 203512US77

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Met Glu Thr Asp Thr Leu Leu Leu Trp Val Leu Leu Leu Trp Val Pro 1 $$ 5 $$ 10 $$ 15

Gly Ser Thr Gly Asp Glu Asp Gln Val Asp Pro Arg Leu 1le Asp Gly $20 \\ 25 \\ 30$

Lys Leu Ser Arg Asp Gln Tyr His Val Leu Phe Asp Ser Tyr Arg Asp $35 \ \ \, 40 \ \ \, 45$

Asn Ile Ala Gly Lys Ser Phe Gln Asn Arg Leu Cys Leu Pro Met Pro 50 60

Ile Asp Val Val Tyr Thr Trp Val Asn Gly Thr Asp Leu Glu Leu Leu 65 70 70 75 80

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Pro Ser Leu Tyr Pro Ser Phe His Ser Ala Ser Asp Ile Phe Asn Val 145 \$150\$

Ala Lys Pro Lys Asn Pro Ser Thr Asn Val Ser Val Val Val Phe Asp $_{165}$ $_{175}$

Ser Thr Lys Asp Val Glu Asp Ala His Ser Gly Leu Leu Lys Gly Asn 180 185

Ser Arg Gln Thr Val Trp Arg Gly Tyr Leu Thr Thr Asp Lys Glu Val \$195\$

Pro Gly Leu Val Leu Met Gln Asp Leu Ala Phe Leu Ser Gly Phe Pro 210 $$\rm 215$$

Pro Thr Phe Lys Glu Thr Asn Gln Leu Lys Thr Lys Leu Pro Glu Asn 225 230 235

Leu Ser Ser Lys Val Lys Leu Leu Gln Leu Tyr Ser Glu Ala Ser Val 245 250 255

Ala Leu Leu Lys Leu Asn Asn Pro Lys Asp Phe Gln Glu Leu Asn Lys $260 \hspace{1.5cm} 265 \hspace{1.5cm} 265 \hspace{1.5cm} 270 \hspace{1.5cm}$

Gln Thr Lys Lys Asn Met Thr Ile Asp Gly Lys Glu Leu Thr Ile Ser $275 \\ 280 \\ 280$

Pro Ala Tyr Leu Leu Trp Asp Leu Ser Ala Ile Ser Gln Ser Lys Gln 290 295 300

Asp Glu Asp Ile Ser Ala Ser Arg Phe Glu Asp Asn Glu Glu Leu Arg 305 310 315

Tyr Ser Leu Arg Ser Ile Glu Arg His Ala Pro Trp Val Arg Asn Ile

325 330 335

Phe Ile Val Thr Asn Gly Gln Ile Pro Ser Trp Leu Asn Leu Asp Asn 340 345 350

Pro Arg Val Thr Ile Val Thr His Gln Asp Val Phe Arg Asn Leu Ser 355 360 365

His Leu Pro Thr Phe Ser Ser Pro Ala Ile Glu Ser His Val His Arg

Ile Glu Gly Leu Ser Gln Lys Phe Ile Tyr Leu Asn Asp Asp Val Met 385 390 395 400

Phe Gly Lys Asp Val Trp Pro Asp Asp Phe Tyr Ser His Ser Lys Gly \$405\$

Gln Lys Val Tyr Leu Thr Trp Pro Val Pro Asn Cys Ala Glu Gly Cys 420 \$420\$

Pro Gly Ser Trp Ile Lys Asp Gly Tyr Cys Asp Lys Ala Cys Asn Asn 435 440 445

Ser Ala Cys Asp Trp Asp Gly Gly Asp Cys Ser Gly Asp Ser Gly Gly 450 455

Ser Arg Tyr Ile Ala Gly Gly Gly Gly Thr Gly Ser Ile Gly Val Gly 465 \$470\$

Gln Pro Trp Gln Phe Gly Gly Gly Ile Asn Ser Val Ser Tyr Cys Asn $485 \hspace{1cm} 490 \hspace{1cm} 495$

Gln Gly Cys Ala Asn Ser Trp Leu Ala Asp Lys Phe Cys Asp Gln Ala 500 505 510

Cys Asn Val Leu Ser Cys Gly Phe Asp Ala Gly Asp Cys Gly Gln Asp 515 520 525

His Phe His Glu Leu Tyr Lys Val Ile Leu Leu Pro Asn Gln Thr His 530 535 540

Val Ala Lys Arg Gly Val Glu Gly Ala Tyr Ser Asp Asn Pro Ile Ile 565 570 575

Arg His Ala Ser Ile Ala Asn Lys Trp Lys Thr Ile His Leu Ile Met $580 \hspace{1.5cm} 585 \hspace{1.5cm} 590 \hspace{1.5cm}$

His Ser Gly Met Asn Ala Thr Thr Ile His Phe Asn Leu Thr Phe Gln 595 $$ 605 $$

Asn Thr Asn Asp Glu Glu Phe Lys Met Gln Ile Thr Val Glu Val Asp 610 620

Thr Arg Glu Gly Pro Lys Leu Asn Ser Thr Ala Gln Lys Gly Tyr Glu 625 630 635 640

Asn Leu Val Ser Pro Ile Thr Leu Leu Pro Glu Ala Glu Ile Leu Phe 645 $\,$ 650 $\,$ 655

Glu Asp Ile Pro Lys Glu Lys Arg Phe Pro Lys Phe Lys Arg His Asp 660 665 670

Val Asn Ser Thr Arg Arg Ala Gln Glu Glu Val Lys Ile Pro Leu Val 675 $\,$ 685 $\,$

Asn Ile Ser Leu Leu Pro Lys Asp Ala Gln Leu Ser Leu Asn Thr Leu $690 \hspace{1.5cm} 695 \hspace{1.5cm} 700 \hspace{1.5cm}$

Asp Leu Gln Leu Glu His Gly Asp Ile Thr Leu Lys Gly Tyr Asn Leu 705 $$ 710 $$ 715 $$ 720

Ser Lys Ser Ala Leu Leu Arg Ser Phe Leu Met As
n Ser Gl
n His Ala 725 730 735

Lys Ile Lys Asn Gln Ala Ile Ile Thr Asp Glu Thr Asn Asp Ser Leu $740 \hspace{1.5cm} 745 \hspace{1.5cm} 750 \hspace{1.5cm}$

Val Ala Pro Gln Glu Lys Gln Val His Lys Ser Ile Leu Pro Asn Ser 755 $\,$ 760 $\,$ 765

Leu Gly Val Ser Glu Arg Leu Gln Arg Leu Thr Phe Pro Ala Val Ser 770 780

Val Lys Val Asn Gly His Asp Gln Gly Gln Asn Pro Pro Leu Asp Leu 785 790 795 800

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Glu Thr Thr Ala Arg Phe Arg Val Glu Thr His Thr Gln Lys Thr Ile $805 \\ 810 \\ 815$

Gly Gly Asn Val Thr Lys Glu Lys Pro Pro Ser Leu Ile Val Pro Leu 820 825 830

Glu Ser Gln Met Thr Lys Glu Lys Lys Ile Thr Gly Lys Glu Lys Glu 835 \$840

Asn Ser Arg Met Glu Glu Asn Ala Glu Asn His Ile Gly Val Thr Glu 850 855 860

Val Leu Leu Gly Arg Lys Leu Gln His Tyr Thr Asp Ser Tyr Leu Gly 865 870 875 880

Phe Leu Pro Trp Glu Lys Lys Lys Tyr Phe Leu Asp Leu Asp Glu 885 890 895

Glu Glu Ser Leu Lys Thr Gln Leu Ala Tyr Phe Thr Asp Ser Lys Asn 900 905 910

Arg Ala Arg Tyr Lys Arg Asp Thr Phe Ala Asp Ser Leu Arg Tyr Val 915 920 925

Asn Lys Ile Leu Asn Ser Lys Phe Gly Phe Thr Ser Arg Lys Val Pro 930 940

Ala His Met Pro His Met Ile Asp Arg Ile Val Met Gln Glu Leu Gln 945 950 955 960

His Ser Glu Asp Met Gln Phe Ala Phe Ser Tyr Phe Tyr Tyr Leu Met 980 \$980

Ser Ala Val Gln Pro Leu Asn Ile Ser Gln Val Phe Asp Glu Val Asp 995 $1000\,$

Thr Asp Gln Ser Gly Val Leu Ser Asp Arg Glu Ile Arg Thr Leu

. . .

Ala Thr Arg Ile His Glu Leu Pro Leu Ser Leu Gln Asp Leu Thr

Gly Leu Glu His Met Leu Ile Asn Cys Ser Lys Met Leu Pro Ala

Asp Ile Thr Gln Leu Asn Asn Ile Pro Pro Thr Gln Glu Ser Tyr

Tyr Asp Pro Asn Leu Pro Pro Val Thr Lys Ser Leu Val Thr Asn

Cys Lys Pro Val Thr Asp Lys Ile His Lys Ala Tyr Lys Asp Lys

Asn Lys Tyr Arg Phe Glu Ile Met Gly Glu Glu Glu Ile Ala Phe

Lys Met Ile Arg Thr Asn Val Ser His Val Val Gly Gln Leu Asp

Asp Ile Arg Lys Asn Pro Arg Lys Phe Val Cys Leu Asn Asp Asn

Ile Asp His Asn His Lys Asp Ala Gln Thr Val Lys Ala Val Leu

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Tyr His Val Leu Phe Asp Ser Tyr Arg Asp Asn Ile Ala Gly Lys Ser 50 55 60

Phe Gln Asn Arg Leu Cys Leu Pro Met Pro Ile Asp Val Val Tyr Thr 65 70 75 80

Trp Val Asn Gly Thr Asp Leu Glu Leu Leu Lys Glu Leu Gln Gln Val

Lys Asn Thr Thr Glu Pro Thr Lys Lys Ser Glu Lys Gln Leu Glu Cys

Leu Leu Thr His Cys Ile Lys Val Pro Met Leu Val Leu Asp Pro Ala 130 135 140

Leu Pro Ala Asn Ile Thr Leu Lys Asp Val Pro Ser Leu Tyr Pro Ser 145 150 155 160

Phe His Ser Ala Ser Asp Ile Phe Asn Val Ala Lys Pro Lys Asn Pro 165 170 175

Ser Thr Asn Val Ser Val Val Val Phe Asp Ser Thr Lys Asp Val Glu

Asp Ala His Ser Gly Leu Leu Lys Gly Asn Ser Arg Gln Thr Val Trp 195 200 205

Gln Asp Leu Ala Phe Leu Ser Gly Phe Pro Pro Thr Phe Lys Glu Thr 225 230 235 240

Asn Gln Leu Lys Thr Lys Leu Pro Glu Asn Leu Ser Ser Lys Val Lys 245 250

Leu Leu Gln Leu Tyr Ser Glu Ala Ser Val Ala Leu Leu Lys Leu Asn $260 \hspace{1.5cm} 265 \hspace{1.5cm} 270 \hspace{1.5cm}$

Asn Pro Lys Asp Phe Gln Glu Leu Asn Lys Gln Thr Lys Lys Asn Met 275 280 285

Thr Ile Asp Gly Lys Glu Leu Thr Ile Ser Pro Ala Tyr Leu Leu Trp 290 295 300

Asp Leu Ser Ala Ile Ser Gln Ser Lys Gln Asp Glu Asp Ile Ser Ala 305 310 315

Ser Arg Phe Glu Asp Asp Glu Glu Leu Arg Tyr Ser Leu Arg Ser Ile 325 330 335

Glu Arg His Ala Pro Trp Val Arg Asn Ile Phe Ile Val Thr Asn Gly 340 345 350

Gln Ile Pro Ser Trp Leu Asn Leu Asp Asn Pro Arg Val Thr Ile Val 355 360 365

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Thr His Gln Asp Val Phe Arg Asn Leu Ser His Leu Pro Thr Phe Ser

Ser Pro Ala Ile Glu Ser His Ile His Arg Ile Glu Gly Leu Ser Gln 385 390 395 400

Lys Phe Ile Tyr Leu Asn Asp Asp Val Met Phe Gly Lys Asp Val Trp \$405\$

Pro Asp Asp Phe Tyr Ser His Ser Lys Gly Gln Lys Val Tyr Leu Thr $420 \hspace{1.5cm} 425 \hspace{1.5cm} 430 \hspace{1.5cm}$

Trp Pro Val Pro Asn Cys Ala Glu Gly Cys Pro Gly Ser Trp Ile Lys 435 440 445

Asp Gly Tyr Cys Asp Lys Ala Cys Asn Asn Ser Ala Cys Asp Trp Asp 450 460

Gly Gly Asp Cys Ser Gly Asn Ser Gly Gly Ser Arg Tyr Ile Ala Gly 465 470475475

Gly Gly Gly Thr Gly Ser Ile Gly Val Gly His Pro Trp Gln Phe Gly 485 490 495

Gly Gly Ile Asn Ser Val Ser Tyr Cys Asn Gln Gly Cys Ala Asn Ser $500 \hspace{1cm} 505 \hspace{1cm} 510$

Gly Phe Asp Ala Gly Asp Cys Gly Gln Asp His Phe His Glu Leu Tyr $530 \hspace{1.5cm} 535 \hspace{1.5cm} 540 \hspace{1.5cm}$

Glu Cys Leu Pro Tyr Phe Ser Phe Ala Glu Val Ala Lys Arg Gly Val 565 570 575

Glu Gly Ala Tyr Ser Asp Asn Pro Ile Ile Arg His Ala Ser Ile Ala

580 585 590

Asn Lys Trp Lys Thr Ile His Leu Ile Met His Ser Gly Met Asn Ala 595 600 605

Thr Thr Ile His Phe Asn Leu Thr Phe Gln Asn Thr Asn Asp Glu Glu

Phe Lys Met Gln Ile Thr Val Glu Val Asp Thr Arg Glu Gly Pro Lys 625 630 635

Thr Leu Leu Pro Glu Ala Glu Ile Leu Phe Glu Asp Ile Pro Lys Glu 660 665 $$ 10 670

Lys Arg Phe Pro Lys Phe Lys Arg His Asp Val Asn Ser Thr Arg Arg 675 680 685

Ala Gln Glu Glu Val Lys Ile Pro Leu Val Asn Ile Ser Leu Leu Pro 690 695 700

Lys Asp Ala Gln Leu Ser Leu Asn Thr Leu Asp Leu Gln Leu Glu His 705 710710715715715

Gly Asp Ile Thr Leu Lys Gly Tyr Asn Leu Ser Lys Ser Ala Leu Leu 735 735

Arg Ser Phe Leu Met Asn Ser Gln His Ala Lys Ile Lys Asn Gln Ala $740 \hspace{1.5cm} 745 \hspace{1.5cm} 750 \hspace{1.5cm}$

Ile Ile Thr Asp Glu Thr Asp Asp Ser Leu Val Ala Pro Gln Glu Lys $755 \ \ \, 760 \ \ \, 765$

Gln Val His Lys Ser Ile Leu Pro Asn Ser Leu Gly Val Ser Glu Arg 770 775 780

Leu Gln Arg Leu Thr Phe Pro Ala Val Ser Val Lys Val Asn Gly His 785 790 795 800

Arg Val Glu Thr His Thr Gln Lys Thr Ile Gly Gly Asn Val Thr Lys 820 825 830

Glu Lys Pro Pro Ser Leu Ile Val Pro Leu Glu Ser Gln Met Thr Lys 835 840 845

Glu Lys Lys Ile Thr Gly Lys Glu Lys Glu Asn Ser Arg Met Glu Glu 850 \$850

Asn Ala Glu Asn His Ile Gly Val Thr Glu Val Leu Leu Gly Arg Lys 865 870 875 880

Leu Gln His Tyr Thr Asp Ser Tyr Leu Gly Phe Leu Pro Trp Glu Lys \$85\$

Lys Lys Tyr Phe Gln Asp Leu Leu Asp Glu Glu Glu Ger Leu Lys Thr $900 \hspace{1cm} 905 \hspace{1cm} 910 \hspace{1cm}$

Gln Leu Ala Tyr Phe Thr Asp Ser Lys Asn Thr Gly Arg Gln Leu Lys $915 \hspace{1.5cm} 920 \hspace{1.5cm} 925 \hspace{1.5cm}$

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<211> 320 <212> PRT

<213> Homo sapiens

<400> 5

Asp Thr Phe Ala Asp Ser Leu Arg Tyr Val Asn Lys Ile Leu Asn Ser 1 $$ 5 $$ 10 $$ 15

Lys Phe Gly Phe Thr Ser Arg Lys Val Pro Ala His Met Pro His Met 20 25 30

Phe Asp Lys Thr Ser Phe His Lys Val Arg His Ser Glu Asp Met Gln 50 60

Phe Ala Phe Ser Tyr Phe Tyr Tyr Leu Met Ser Ala Val Gln Pro Leu 65 70 70 80

Asn Ile Ser Gln Val Phe Asp Glu Val Asp Thr Asp Gln Ser Gly Val

Leu Ser Asp Arg Glu Ile Arg Thr Leu Ala Thr Arg Ile His Glu Leu 100 105 110

Pro Leu Ser Leu Gln Asp Leu Thr Gly Leu Glu His Met Leu Ile Asn \$115\$ \$120\$ \$125\$

Cys Ser Lys Met Leu Pro Ala Asp Ile Thr Gln Leu Asn Asn Ile Pro 130 140

Pro Thr Glu Ser Tyr Tyr Asp Pro Asn Leu Pro Pro Val Thr Lys 145 $$ 150 $$ 150 $$ 155 $$ 160

Ser Leu Val Thr Asn Cys Lys Pro Val Thr Asp Lys Ile His Lys Ala 165 170 175

Tyr Lys Asp Lys Asn Lys Tyr Arg Phe Glu Ile Met Gly Glu Glu Glu 180 \$180\$

Ile Ala Phe Lys Met Ile Arg Thr Asn Val Ser His Val Val Gly Gln
195 200 205

Leu Asp Asp Ile Arg Lys Asn Pro Arg Lys Phe Val Cys Leu Asn Asp 210 215 220

Asn Ile Asp His Asn His Lys Asp Ala Gln Thr Val Lys Ala Val Leu 225 230 235 240

Arg Asp Phe Tyr Glu Ser Met Phe Pro Ile Pro Ser Gln Phe Glu Leu 245 250 255

Pro Arg Glu Tyr Arg Asn Arg Phe Leu His Met His Glu Leu Glu Glu 260 265 270

Trp Arg Ala Tyr Arg Asp Lys Leu Lys Phe Trp Thr His Cys Val Leu 275 280 285

Ala Thr Leu Ile Met Phe Thr Ile Phe Ser Phe Phe Ala Glu Gln Leu 290 295 300

Ile Ala Leu Lys Arg Lys Ile Phe Pro Arg Arg Ile His Lys Glu

60

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240

300

360

420

540

600 660

720 780

840

900

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<211> 1219 DNA

<213> Homo sapiens

<400> 6

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- <211> 305 <212> PRT <213> Homo sapiens
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- Gly Gly Pro Ala Pro Ala Gly Ala Ala Lys Met Lys Val Val Glu Glu 20
- Pro Asn Ala Phe Gly Val Asn Asn Pro Phe Leu Pro Gln Ala Ser Arg
- Leu Gln Ala Lys Arg Asp Pro Ser Pro Val Ser Gly Pro Val His Leu
- Phe Arg Leu Ser Gly Lys Cys Phe Ser Leu Val Glu Ser Thr Tyr Lys
- Tyr Glu Phe Cys Pro Phe His Asn Val Thr Gln His Glu Gln Thr Phe
- Arg Trp Asn Ala Tyr Ser Gly Ile Leu Gly Ile Trp His Glu Trp Glu 105 100
- Ile Ala Asn Asn Thr Phe Thr Gly Met Trp Met Arg Asp Gly Asp Ala 120 125
- Cys Arg Ser Arg Ser Arg Gln Ser Lys Val Glu Leu Ala Cys Gly Lys 135 130
- Ser Asn Arg Leu Ala His Val Ser Glu Pro Ser Thr Cys Val Tyr Ala 150 155 145
- Leu Thr Phe Glu Thr Pro Leu Val Cys His Pro His Ala Leu Leu Val 165
- Tyr Pro Thr Leu Pro Glu Ala Leu Gln Arg Gln Trp Asp Gln Val Glu 180
- Gln Asp Leu Ala Asp Glu Leu Ile Thr Pro Gln Gly His Glu Lys Leu 200 205 195

Leu Arg Thr Leu Phe Glu Asp Ala Gly Tyr Leu Lys Thr Pro Glu Glu 210 Asn Glu Pro Thr Gln Leu Glu Gly Gly Pro Asp Ser Leu Gly Phe Glu 225 230 Thr Leu Glu Asn Cys Arg Lys Ala His Lys Glu Leu Ser Lys Glu Ile 245 Lys Arg Leu Lys Gly Leu Leu Thr Gln His Gly Ile Pro Tyr Thr Arg 260 265 Pro Thr Glu Thr Ser Asn Leu Glu His Leu Gly His Glu Thr Pro Arg 275 Ala Lys Ser Pro Glu Gln Leu Arg Gly Asp Pro Gly Leu Arg Gly Ser 290 Leu 305 <210> <211> 5229 <212> DNA <213> Mus musculus <400> 8 ggcggtgaag gggtgatgct gttcaagctc ctgcagagac agacctatac ctgcctatcc 60 cacaggtatg ggetetacgt etgettegtg ggegtegttg teaccategt eteggettte 120 caqttcqqaq aqqtqqttct ggaatggagc cgagatcagt accatgtttt gtttgattcc 180 tacaqaqaca acattgctgg gaaatccttt cagaatcggc tctgtctgcc catgccaatc 240 gacgtggttt acacctgggt gaatggcact gaccttgaac tgctaaagga gctacagcag 300 360 qtccqaqaqc acatggagga agagcagaga gccatgcggg aaaccctcgg gaagaacaca 420 accgaaccga caaagaagag tgagaagcag ctggaatgtc tgctgacgca ctgcattaag gtgcccatgc ttgttctgga cccggccctg ccagccacca tcaccctgaa ggatctgcca 480

540

600

660

accetttace catettteca egegtecage gacatgttea atgttgcgaa accaaaaaat

cegtetacaa atgteecegt tgtegttttt gacactacta aggatgttga agacgeecat

qetqqaeeqt ttaagggagg eeagcaaaca gatgtttgga gagcetactt gacaacagac

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aaaaaaaaa

Arg Tyr Gly Leu Tyr Val Cys Phe Val Gly Val Val Val Thr Ile Val 20 25 30

Ser Ala Phe Gln Phe Gly Glu Val Val Leu Glu Trp Ser Arg Asp Gln

5229

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<212> PRT

<213> Mus musculus

<400> 9

35 40 45

Tyr His Val Leu Phe Asp Ser Tyr Arg Asp Asn Ile Ala Gly Lys Ser

Phe Gln Asn Arg Leu Cys Leu Pro Met Pro Ile Asp Val Val Tyr Thr 65 70 75 80

Trp Val Asn Gly Thr Asp Leu Glu Leu Leu Lys Glu Leu Gln Gln Val 85 90 95

Arg Glu His Met Glu Glu Glu Gln Arg Ala Met Arg Glu Thr Leu Gly 100 $$105\$

Lys Asn Thr Thr Glu Pro Thr Lys Lys Ser Glu Lys Gln Leu Glu Cys 115 120 125

Leu Leu Thr His Cys Ile Lys Val Pro Met Leu Val Leu Asp Pro Ala 130 $$135\$

Leu Pro Ala Thr Ile Thr Leu Lys Asp Leu Pro Thr Leu Tyr Pro Ser 145 150 155 160

Phe His Ala Ser Ser Asp Met Phe Asn Val Ala Lys Pro Lys Asn Pro $165 \\ 170 \\ 175$

Ser Thr Asn Val Pro Val Val Val Phe Asp Thr Thr Lys Asp Val Glu
180 185 190

Asp Ala His Ala Gly Pro Phe Lys Gly Gly Gln Gln Thr Asp Val Trp $195 \hspace{0.5cm} 200 \hspace{0.5cm} 205 \hspace{0.5cm}$

Arg Ala Tyr Leu Thr Thr Asp Lys Asp Ala Pro Gly Leu Val Leu Ile 210 225 220

Gln Gly Leu Ala Phe Leu Ser Gly Phe Pro Pro Thr Phe Lys Glu Thr 225 $$ 230 $$ 235 $$ 240

Ser Gln Leu Lys Thr Lys Leu Pro Arg Lys Ala Phe Pro Leu Lys Ile $_{245}$ $_{250}$ $_{25}$

Lys Leu Leu Arg Leu Tyr Ser Glu Ala Ser Val Ala Leu Leu Lys Leu 260 265 270

Asn Asn Pro Lys Gly Phe Gln Glu Leu Asn Lys Gln Thr Lys Lys Asn 275 280 285

* *

Met Thr Ile Asp Gly Lys Glu Leu Thr Ile Ser Pro Ala Tyr Leu Leu 290 295 300

Trp Asp Leu Ser Ala Ile Ser Gln Ser Lys Gln Asp Glu Asp Ala Ser 305 $$ 310 $$ 315 $$ 320

Ile Glu Arg His Ala Pro Trp Val Arg Asn Ile Phe Ile Val Thr Asn \$340\$ \$345\$ \$350

Gly Gln Ile Pro Ser Trp Leu Asn Leu Asp Asn Pro Arg Val Thr Ile 355 360 365

Ser Ser Pro Ala Ile Glu Ser His Ile His Arg Ile Glu Gly Leu Ser 385 390 395 400

Gln Lys Phe Ile Tyr Leu Asn Asp Asp Val Met Phe Gly Lys Asp Val 415

Trp Pro Asp Asp Phe Tyr Ser His Ser Lys Gly Gln Lys Val Tyr Leu $420 \hspace{1.5cm} 425 \hspace{1.5cm} 430 \hspace{1.5cm}$

Thr Trp Pro Val Pro Asn Cys Ala Glu Gly Cys Pro Gly Ser Trp Ile 435 440 445

Lys Asp Gly Tyr Cys Asp Lys Ala Cys Asn Thr Ser Pro Cys Asp Trp 450 455 460

Asp Gly Gly Asn Cys Ser Gly Asn Thr Ala Gly Asn Arg Phe Val Ala 465 470475475

Arg Gly Gly Gly Thr Gly Asn Ile Gly Ala Gly Gln His Trp Gln Phe 485 490 495

- Ser Trp Leu Ala Asp Lys Phe Cys Asp Gln Ala Cys Asn Val Leu Ser 515 520 525
- Cys Gly Phe Asp Ala Gly Asp Cys Gly Gln Asp His Phe His Glu Leu 530 535 540
- Tyr Lys Val Thr Leu Leu Pro Asn Gln Thr His Tyr Val Val Pro Lys 545 550 555 560
- Gly Glu Tyr Leu Ser Tyr Phe Ser Phe Ala Asn Ile Ala Arg Lys Arg $565 \hspace{1cm} 570 \hspace{1cm} 575$
- Ile Glu Gly Thr Tyr Ser Asp Asn Pro Ile Ile Arg His Ala Ser Ile 580 585 590
- Ala Asn Lys Trp Lys Thr Leu His Leu Ile Met Pro Gly Gly Met Asn 595 600 605
 - Ala Thr Thr Ile Tyr Phe Asn Leu Thr Leu Gln Asn Ala Asn Asp Glu $_{610}$ $_{620}$
- Glu Phe Lys Ile Gln Ile Ala Val Glu Val Asp Thr Arg Glu Ala Pro 625 $$ 630 $$ 635 $$ 640
- Lys Leu Asn Ser Thr Thr Gln Lys Ala Tyr Glu Ser Leu Val Ser Pro $645 \\ 650 \\ 655$
- Val Thr Pro Leu Pro Gln Ala Asp Val Pro Phe Glu Asp Val Pro Lys 660 665 670
- Glu Lys Arg Phe Pro Lys Ile Arg Arg His Asp Val Asn Ala Thr Gly 675 680 685
- Arg Phe Gln Glu Glu Val Lys Ile Pro Arg Val Asn Ile Ser Leu Leu 690 700
- Pro Lys Glu Ala Gln Val Arg Leu Ser Asn Leu Asp Leu Gln Leu Glu 705 710 715 720

Arg Gly Asp Ile Thr Leu Lys Gly Tyr Asn Leu Ser Lys Ser Ala Leu 725 730 735

Leu Arg Ser Phe Leu Gly Asn Ser Leu Asp Thr Lys Ile Lys Pro Gln 740 745 750

Ala Arg Thr Asp Glu Thr Lys Gly Asn Leu Glu Val Pro Gln Glu Asn 755 $$ 760 $$ 765

Pro Ser His Arg Arg Pro His Gly Phe Ala Gly Glu His Arg Ser Glu 770 775 780

Arg Trp Thr Ala Pro Ala Glu Thr Val Thr Val Lys Gly Arg Asp His 785 790795795

Ala Leu Asn Pro Pro Pro Val Leu Glu Thr Asn Ala Arg Leu Ala Gln 805 810 815

Pro Thr Leu Gly Val Thr Val Ser Lys Glu Asn Leu Ser Pro Leu Ile 820 825 830

Val Pro Pro Glu Ser His Leu Pro Lys Glu Glu Glu Ser Asp Arg Ala 835 840 845

Glu Gly Asn Ala Val Pro Val Lys Glu Leu Val Pro Gly Arg Arg Leu 850 855 860

Gln Gln Asn Tyr Pro Gly Phe Leu Pro Trp Glu Lys Lys Lys Tyr Phe 865 870 875 880

Gln Asp Leu Leu Asp Glu Glu Glu Ser Leu Lys Thr Gln Leu Ala Tyr 885 890 895

Phe Thr Asp Arg Lys His Thr Gly Arg Gln Leu Lys

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<211> 328

<212> PRT <213> Mus musculus

<400> 10

Asp Thr Phe Ala Asp Ser Leu Arg Tyr Val Asn Lys Ile Leu Asn Ser 1 $$ 5 $$ 10 $$ 15

Lys Phe Gly Phe Thr Ser Arg Lys Val Pro Ala His Met Pro His Met

Ile Asp Arg Ile Val Met Gln Glu Leu Gln Asp Met Phe Pro Glu Glu 35 40 45

Phe Asp Lys Thr Ser Phe His Lys Val Arg His Ser Glu Asp Met Gln 50 60

Phe Ala Phe Ser Tyr Phe Tyr Tyr Leu Met Ser Ala Val Gln Pro Leu 65 70 75 80

Asn Ile Ser Gln Val Phe His Glu Val Asp Thr Asp Gln Ser Gly Val 85 90 95

Leu Ser Asp Arg Glu Ile Arg Thr Leu Ala Thr Arg Ile His Asp Leu 100 105 110

Pro Leu Ser Leu Gln Asp Leu Thr Gly Leu Glu His Met Leu Ile Asn \$115\$

Cys Ser Lys Met Leu Pro Ala Asn Ile Thr Gln Leu Asn Asn Ile Pro $130\,$

Pro Thr Gln Glu Ala Tyr Tyr Asp Pro Asn Leu Pro Pro Val Thr Lys 145 \$150\$

Ser Leu Val Thr Asn Cys Lys Pro Val Thr Asp Lys Ile His Lys Ala 165 170 175

Tyr Lys Asp Lys Asn Lys Tyr Arg Phe Glu Ile Met Gly Glu Glu Glu 180 185 190

Ile Ala Phe Lys Met Ile Arg Thr Asn Val Ser His Val Val Gly Gln 195 \$200\$

Leu Asp Asp Ile Arg Lys Asn Pro Arg Lys Phe Val Cys Leu Asn Asp 210 215

Asn Ile Asp His Asn His Lys Asp Ala Arg Thr Val Lys Ala Val Leu 225 230 235 240

i	Arg	Asp	Phe	Tyr	Glu 245	Ser	Met	Phe	Pro	Ile 250	Pro	Ser	Gln	Phe	Glu 255	Leu		
	Pro	Arg	Glu	Tyr 260	Arg	Asn	Arg	Phe	Leu 265	His	Met	His	Glu	Leu 270	Gln	Glu		
	Trp	Arg	Ala 275	Tyr	Arg	Asp	Lys	Leu 280	Lys	Phe	Trp	Thr	His 285	Сув	Val	Leu		
	Ala	Thr 290	Leu	Ile	Ile	Phe	Thr 295	Ile	Phe	Ser	Phe	Phe 300	Ala	Glu	Gln	Ile		
	Ile 305	Ala	Leu	Lys	Arg	Lys 310	Ile	Phe	Pro	Arg	Arg 315	Arg	Ile	His	Lys	Glu 320		
	Ala	Ser	Pro	Asp	Arg 325	Ile	Arg	Val										
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		1 > 2 >	(186	_fea)(a,	186)		g											
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	cat	tcgg	gtg	agcg	gato	ac g	gtcc	tgcg	g ct	tggg	gacc	gag	cate	gct	ggtt	cttctg	18	0
	acc	ttnt	caa	ttcc	atag	gc t	gaat	aacc	c gt	tctt	gccc	cag	gcaa	gcc	gcct	tcagcc	24	١0
	caa	gaga	.gag	cctt	cago	tg t	atco	cgca	a at	taag	agaa	att	aatt	tca	aacg	atttag	30	0
	aaa	gtat	tet	agco	aggo	ga t	gatg	gege	a co	cctt	taat	ccc	agca	ctt	ggga	ggcaga	36	50
	ggc	aggo	aga	tttc	cgaç	rtt c	aagg	jccat	c ag	aact	gact	gta	cato	tta	gtac	agttta	42	2 C
	gca	tgtg	atc	agag	gatet	ga a	tcac	aaag	c to	ggco	tgcg	tgg	taaa	ıgca	ggto	ctttct	48	3 C
	aat	aagg	ttg	cagt	ttac	at t	ttct	ttct	t aa	ctct	ttta	tto	ttte	gaga	cagg	gtttct	54	ı C

caacagtggg tgtcctggaa ctcacttttg taaaccaggc tgcccttaaa ctcacaaagc 600

tetgteagee tetgeeteet gagtgetggg attaaaggte cacaccetgt teatteattt 660 ttaatttttg agactgggtc tcattatgtg gccctagaca gatactgaga gcctcctcca 720 caggaacaag catgggaatc ctgccacaga caaccagttc tgtggtctgg agatgagttt 780 gtcagtccct aggagttagg tcagcctgcc tctgcattcc caataattta ggaaaggagc 840 ttggggegtt etggeettga tggttagtge ceteetgeea acettagett eeagetttag 900 qqqtaqcaqa gtttataccq atgctaaact gctgttgtgt tcttccccag ggcccctgca 960 totottcaga cttgctggca agtgctttag cctagtggag tccacgtgag tgccaggctg 1020 gtgggtggag tgggcggagt ctgcagagct cctgatgtgc ctgtgtttcc caggtacaag 1080 tatgaattet geeettteea caacgteace cageacgage agacetteeg etggaatgee 1140 tacageggga teettggcat etggcatgag tgggaaatca tcaacaatac ettcaaggge 1200 atgtggatga ctgatgggga ctcctgccac tcccggagcc ggcagagcaa ggtggagctc 1260 acctgtggaa agatcaaccg actggcccac gtgtctgagc caagcacctg tgtctatgca 1320 ttgacattcg agacccctct tgtttgccat ccccactctt tgttagtgta tccaactctg 1380 toagaageec tgeageagee ettggaceag gtggaacagg acetggeaga tgaactgate 1440 acaccacagg gctatgagaa gttgctaagg gtactttttg aggatgctgg ctacttaaag 1500 qtcccaqgag aaacccatcc cacccagetg gcaggaggtt ccaagggcct ggggettgag 1560 actotggaca actgtagaaa ggcacatgca gagetgtcac aggaggtaca aagactgacg 1620 agtotgotgo aacagcatgo aatoccocac actoagocca caggicagto tgoctgocct 1680 qqtcaqctgc cagccactcc ggggcctgca gcactggggc agatctttat tgctacccat 1740 tetqqcagaa accaetcaet etcagcaect gggtcagcag etceccatag gtgcaatcgc 1800 aqcaqaqcat etgeggagtg acccaggact acgtgggaac atcctgtgag caaggtggcc 1860 acqaaqaata gaaatateet gagetttgag tgteetttea cagagtgaac aaaactggtg 1920 tqqtqtaqac acqgcttctt ttggcatatt ctagatcaga cagtgtcact gacaaacaag 1980 agggacetge tggccageet ttgttgtgcc caaagatcca gacaaaataa agattcaaag 2040 2070 ttttaattaa aaaaaaaaa aaaggaattc

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<211> 307

<212> PRT

<213> Mus musculus

<400> 12

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Gln Gly Pro Ala Pro Ala Cys Ala Gly Lys Met Lys Val Val Glu Glu 20 25 30

Pro Asn Thr Phe Gly Leu Asn Asn Pro Phe Leu Pro Gln Ala Ser Arg 35 40 45

Leu Gln Pro Lys Arg Glu Pro Ser Ala Val Ser Gly Pro Leu His Leu 50 60

Phe Arg Leu Ala Gly Lys Cys Phe Ser Leu Val Glu Ser Thr Tyr Lys 65 70 75 80

Tyr Glu Phe Cys Pro Phe His Asn Val Thr Gln His Glu Gln Thr Phe 85 90 90 95

Arg Trp Asn Ala Tyr Ser Gly Ile Leu Gly Ile Trp His Glu Trp Glu $100 \hspace{1cm} 105 \hspace{1cm} 110 \hspace{1cm}$

Ile Ile Asn Asn Thr Phe Lys Gly Met Trp Met Thr Asp Gly Asp Ser 115 120 125

Cys His Ser Arg Ser Arg Gln Ser Lys Val Glu Leu Thr Cys Gly Lys $130 \ \ 135 \ \ 140$

Leu Thr Phe Glu Thr Pro Leu Val Cys His Pro His Ser Leu Leu Val 165 170 175

Tyr Pro Thr Leu Ser Glu Ala Leu Gln Gln Arg Leu Asp Gln Val Glu $180\,$

Gln Asp Leu Ala Asp Glu Leu Ile Thr Pro Gln Gly Tyr Glu Lys Leu 195 200 205

Leu Arg Val Leu Phe Glu Asp Ala Gly Tyr Leu Lys Val Pro Gly Glu 210 215 220

Thr His Pro Thr Gln Leu Ala Gly Gly Ser Lys Gly Leu Gly Leu Glu 225 230 230 235	
Thr Leu Asp Asn Cys Arg Lys Ala His Ala Glu Leu Ser Gln Glu Val 245 250 255	
Gln Arg Leu Thr Ser Leu Leu Gln Gln His Gly Ile Pro His Thr Gln 260 265 270	
Pro Thr Glu Thr Thr His Ser Gln His Leu Gly Gln Gln Leu Pro Ile 275 280 285	
Gly Ala Ile Ala Ala Glu His Leu Arg Ser Asp Pro Gly Leu Arg Gly 290 295 300	
Asn Ile Leu 305	
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taaccccaaa ggtttccccg agctgaacaa gcagaccaag aagaacatga gcatcagtgg	180
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gaaggaactg gccatcagce ctgcctattt gttgtgggac ctgaggcca tcagccagec	300
actgagatct ategagagac atgattecat gagteettta tgaattetgg ccatatette	360
aatcatgatc toagtagtat toototgaaa tggcacacat tittotaatg agaacttgaa	420
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Glu Asn Leu Ser Ser Lys Ile Lys Leu Leu Gln Leu Tyr Ser Glu Ala \$20\$ Ser Val Ala Leu Leu Lys Leu Asn Asn Pro Lys Gly Phe Pro Glu Leu

35 40 45

Asn Lys Gln Thr Lys Lys Asn Met Ser Ile Ser Gly Lys Glu Leu Ala 50 55 60

Ile Ser Pro Ala Tyr Leu Leu Trp Asp Leu Ser Ala Ile Ser Gln Ser 65 70 70 75 80

Lys Gln Asp Glu Asp Val Ser Ala Ser Arg Phe Glu Asp Asn Glu Glu 85 90 95

Leu Arg Tyr Ser Leu Arg Ser Ile Glu Arg His Asp Ser Met Ser Pro $100 \\ 105 \\ 110$

Leu

2210 15
2211 1105
2212 DNA
2213 Drosophila melanogaster
2220 misc_feature
2222 (903) . (903)
2233 n is a, g, t, or c

2200
2212 misc_feature
2222 (935) . (935)
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ggatccagaa gtgcgaaccn tttta

gtnatgagtg tagangaaat cttcgatgaa gtcgacaccg gacggtttgg ncacctggtc

1080

1105

<210> 16 <211> 502

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Ser Leu Glu Lys His Ala Ala Trp Ile Arg His Val Tyr Ile Val Thr

As Gly Gln Ile Pro Ser Trp Leu Asp Leu Ser Tyr Glu Arg Val Thr 35 40 45

Val Val Pro His Glu Val Leu Ala Pro Asp Pro Asp Gln Leu Pro Thr 50 60

Ser Lys Arg Phe Leu Tyr Leu Asn Asp Asp Ile Phe Leu Gly Ala Pro $85 \hspace{0.25in} 90 \hspace{0.25in} 95$

Leu Tyr Pro Glu Asp Leu Tyr Thr Glu Ala Glu Gly Val Arg Val Tyr 100 105 110

Gln Ala Trp Met Val Pro Gly Cys Ala Leu Asp Cys Pro Trp Thr Tyr 115 120 125

Ile Gly Asp Gly Ala Cys Asp Arg His Cys Asn Ile Asp Ala Cys Gln 130 135 140

Phe Asp Gly Gly Asp Cys Ser Glu Thr Gly Pro Ala Ser Asp Ala His 145 \$150\$

Val Ile Pro Pro Ser Lys Glu Val Leu Glu Val Gln Pro Ala Ala Val
165 170 175

Pro Gln Ser Arg Val His Arg Phe Pro Gln Met Gly Leu Gln Lys Leu 180 185 190

Phe Arg Arg Ser Ser Ala Asn Phe Lys Asp Val Met Arg His Arg Asn 195 200 205

Val Ser Thr Leu Lys Glu Leu Arg Arg Ile Val Glu Arg Phe Asn Lys 210 215 220

Ala Lys Leu Met Ser Leu Asn Pro Glu Leu Glu Thr Ser Ser Ser Glu 225 230 235

Pro Gln Thr Thr Gln Arg His Gly Leu Arg Lys Glu Asp Phe Lys Ser 245 250 255

Ser Thr Asp Ile Tyr Ser His Ser Leu Ile Ala Thr Asn Met Leu Leu $_{\rm 260}$ $_{\rm 265}$

Asn Arg Ala Tyr Gly Phe Lys Ala Arg His Val Leu Ala His Val Gly $275 \hspace{1cm} 280 \hspace{1cm} 285 \hspace{1cm}$

Phe Leu Ile Asp Lys Asp Ile Val Glu Ala Met Gln Arg Arg Phe His 290 295 300

Gln Gln Ile Leu Asp Thr Ala His Gln Arg Phe Arg Ala Pro Thr Asp 305 \$310\$ \$315

Leu Gln Tyr Ala Phe Ala Tyr Tyr Ser Phe Leu Met Ser Glu Thr Lys 325 330 335

Ala Thr Trp Ser Asp Arg Glu Val Arg Thr Phe Leu Thr Arg Ile Tyr \$355\$ \$360\$

Gln Pro Pro Leu Asp Trp Ser Ala Met Arg Tyr Phe Glu Glu Val Val 370 375 380

Gln Asn Cys Thr Arg Asn Leu Gly Met His Leu Lys Val Asp Thr Val $_{385}$ $_{390}$ $_{395}$ $_{400}$

Glu His Ser Thr Leu Val Tyr Glu Arg Tyr Glu Asp Ser Asn Leu Pro $405 \hspace{0.25in} 410 \hspace{0.25in} 410 \hspace{0.25in} 415$

Thr Ile Thr Arg Asp Leu Val Val Arg Cys Pro Leu Leu Ala Glu Ala 420 425 430

Leu Ala Asn Phe Ala Val Arg Pro Lys Tyr Asn Phe His Val Ser 435 440 445

Pro Lys Arg Thr Ser His Ser Asn Phe Met Met Leu Thr Ser Asn Leu 450 455 460

Thr Glu Val Val Glu Ser Leu Asp Arg Leu Arg Arg Asn Pro Arg Lys 465 470 475 480

Phe Asn Cys Ile Asn Asp Asn Leu Asp Ala Asn Arg Gly Glu Asp Asn 485 490 495

atggegacot ccaegggteg etggettete etceggettg caetattegg ettectetgg

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Glu Asp Gly Ala Pro Ser

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<212> DNA

<213> Homo sapiens

<400> 17

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<211> 515
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<400> 18

Met Ala Thr Ser Thr Gly Arg Trp Leu Leu Leu Arg Leu Ala Leu Phe 1 $$ 5 $$ 10 $$ 15

Gly Phe Leu Trp Glu Ala Ser Gly Gly Leu Asp Ser Gly Ala Ser Arg $20 \hspace{1cm} 25 \hspace{1cm} 30$

Asp Asp Asp Leu Leu Pro Tyr Pro Arg Ala Arg Ala Arg Leu Pro

<212> PRT

<213> Homo sapiens

35 40 45

Arg Asp Cys Thr Arg Val Arg Ala Gly Asn Arg Glu His Glu Ser Trp 50 55 60

Pro Pro Pro Pro Ala Thr Pro Gly Ala Gly Gly Leu Ala Val Arg Thr 65 70 75 80

Phe Val Ser His Phe Arg Asp Arg Ala Val Ala Gly His Leu Thr Arg 85 9095

Ala Val Glu Pro Leu Arg Thr Phe Ser Val Leu Glu Pro Gly Gly Pro
100 105 110

Gly Gly Cys Ala Ala Arg Arg Arg Ala Thr Val Glu Glu Thr Ala Arg 115 \$120\$

Ala Ala Asp Cys Arg Val Ala Gln Asn Gly Gly Phe Phe Arg Met Asn 130 $$135\$

Ser Gly Glu Cys Leu Gly Asn Val Val Ser Asp Glu Arg Arg Val Ser 145 $$ 150 $$ $$ 155 $$ $$ 160

Ser Ser Gly Gly Leu Gln Asn Ala Gln Phe Gly Ile Arg Arg Asp Gly 165 \$170\$

Thr Leu Val Thr Gly Tyr Leu Ser Glu Glu Glu Val Leu Asp Thr Glu 180 $$185\$

Asn Pro Phe Val Gln Leu Leu Ser Gly Val Val Trp Leu Ile Arg Asn 195 200200205

Gly Ser Ile Tyr Ile Asn Glu Ser Gln Ala Thr Glu Cys Asp Glu Thr 210 215 220

Gln Glu Thr Gly Ser Phe Ser Lys Phe Val Asn Val Ile Ser Ala Arg 225 230235

Thr Ala Ile Gly His Asp Arg Lys Gly Gln Leu Val Leu Phe His Ala 245 250 255

Asp Gly His Thr Glu Gln Arg Gly Ile Asn Leu Trp Glu Met Ala Glu 260 265 270

- Phe Leu Leu Lys Gln Asp Val Val Asn Ala Ile Asn Leu Asp Gly Gly 275 280 285
- Gly Ser Ala Thr Phe Val Leu Asn Gly Thr Leu Ala Ser Tyr Pro Ser 290 \$295\$
- Asp His Cys Gln Asp Asn Met Trp Arg Cys Pro Arg Gln Val Ser Thr 305 $$ 310 $$ 315 $$ 320
- Val Val Cys Val His Glu Pro Arg Cys Gln Pro Pro Asp Cys His Gly 325 330 335
- His Gly Thr Cys Val Asp Gly His Cys Gln Cys Thr Gly His Phe Trp 340 345 350
- Gln His Gly Leu Cys Thr Glu Thr Gly Cys Arg Cys Asp Ala Gly Trp $370 \\ \hspace*{1.5cm} 375 \\ \hspace*{1.5cm} 380 \\ \hspace*{1.5cm}$
- Thr Gly Ser Asn Cys Ser Glu Glu Cys Pro Leu Gly Trp His Gly Pro 385 390395
- Lys Thr Gly Asn Cys Ser Val Ser Arg Val Lys Gln Cys Leu Gln Pro $420 \ \ \, 425 \ \ \, 430$
- Pro Glu Ala Thr Leu Arg Ala Gly Glu Leu Ser Phe Phe Thr Arg Thr 435 \$440\$
- Ile Ala Ala Asn Leu Ser Leu Leu Leu Ser Arg Ala Glu Arg Asn Arg 465 470 475 480
- Arg Leu His Gly Asp Tyr Ala Tyr His Pro Leu Gln Glu Met Asn Gly 485 490 495

Glu Pro Leu Ala Ala Glu Lys Glu Gln Pro Gly Gly Ala His Asn Pro 500 505 510

Phe Lys Asp 515

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1260

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<211> 492 <212> PRT

<213> Mus musculus

<400> 20

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Arg Pro Ser Arg Asp Cys Ala Arg Val Arg Ser Gly Ser Pro Glu Gln 20 25 30

Glu Ser Trp Pro Pro Pro Pro Leu Ala Thr His Glu Pro Arg Ala Pro 35 40 45

Ser His His Ala Ala Val Arg Thr Phe Val Ser His Phe Glu Gly Arg 50 $\,$ 55 $\,$ 60 $\,$

Ala Val Ala Gly His Leu Thr Arg Val Ala Asp Pro Leu Arg Thr Phe 65 $$ 70 $$ 70 $$ 75 $$ 80

Ser Val Leu Glu Pro Gly Gly Ala Gly Gly Cys Gly Gly Arg Ser Ala 85 90 90 95

- Ala Ala Thr Val Glu Asp Thr Ala Val Arg Ala Gly Cys Arg Ile Ala 100 \$105\$
- Gln Asn Gly Gly Phe Phe Arg Met Ser Thr Gly Glu Cys Leu Gly Asn 115 \$120\$
- Val Val Ser Asp Gly Arg Leu Val Ser Ser Ser Gly Gly Leu Gln Asn 130 135 140
- Ala Gln Phe Gly Ile Arg Arg Asp Gly Thr Ile Val Thr Gly Ser Cys 145 150 150
- Leu Glu Glu Val Leu Asp Pro Val Asp Pro Phe Val Gln Leu Leu 165 \$170\$
- Ser Gly Val Val Trp Leu Ile Arg Asn Gly Asn Ile Tyr Ile Asn Glu 180 185 190
- Ser Gln Ala Ile Glu Cys Asp Glu Thr Gln Glu Thr Gly Ser Phe Ser 195 \$200\$
- Lys Phe Val Asn Val Met Ser Ala Arg Thr Ala Val Gly His Asp Arg 210 220
- Glu Gly Gln Leu Ile Leu Phe His Ala Asp Gly Gln Thr Glu Gln Arg 225 230 235 240
- Gly Leu Asn Leu Trp Glu Met Ala Glu Phe Leu Arg Gln Gln Asp Val 245 250 255
- Val Asn Ala Ile Asn Leu Asp Gly Gly Gly Ser Ala Thr Phe Val Leu $260 \hspace{1cm} 265 \hspace{1cm} 270 \hspace{1cm}$
- Asn Gly Thr Leu Ala Ser Tyr Pro Ser Asp His Cys Gln Asp Asn Met 275 \$280\$
- Trp Arg Cys Pro Arg Gln Val Ser Thr Val Val Cys Val His Glu Pro 290 295 300
- Arg Cys Gln Pro Pro Asp Cys Ser Gly His Gly Thr Cys Val Asp Gly 305 310 315

His Cys	Glu Cy	s Thr 325	Ser	His	Phe	Trp	Arg 330	Gly	Glu	Ala	Cys	Ser 335	Glu
Leu Asp	Cys Gl 34		Ser	Asn	Cys	Ser 345	Gln	His	Gly	Leu	Суs 350	Thr	Ala
Gly Cys	His Cy 355	s Asp	Ala	Gly	Trp 360	Thr	Gly	Ser	Asn	Cys 365	Ser	Glu	Glu
Cys Pro 370	Leu Gl	y Trp	Tyr	Gly 375		Gly	Cys	Gln	Arg 380	Pro	Cys	Gln	Cys
Glu His 385	Gln Cy	3 Phe	Cys 390	Asp	Pro	Gln	Thr	Gly 395	Asn	Cys	Ser	Ile	Ser 400
Gln Val	Arg Gl:	n Cys 405	Leu	Gln	Pro	Thr	Glu 410	Ala	Thr	Pro	Arg	Ala 415	Gly
Glu Leu	Ala Se 42	Phe	Thr	Arg	Thr	Thr 425	Trp	Leu	Ala	Leu	Thr 430	Leu	Thr
Leu Ile	Phe Le	ı Leu	Leu	Ile	Ser 440	Thr	Gly	Val	Asn	Val 445	Ser	Leu	Phe
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